

# FIG. 1A

## KCNQ5 cDNA coding sequence

atgaaggatg tggagtcggg ccggggcagg gtgctgctga 40  
 actcggcagc cgccaggggc gacggcctgc tactgctggg 80  
 caccgcgcgc gccacgcttg gtggcggcgg cggtggcctg 120  
 agggagagcc gccggggcaa gcagggggcc cggatgagcc 160  
 tgctggggaa gccgctctct tacacgagta gccagagctg 200  
 ccggcgcaac gtcaagtacc ggcggttgca gaactacctg 240  
 tacaacgtgc tggagagacc ccgcggctgg gcgttcatct 280  
 accacgcttt cgtttttctc cttgtctttg gttgcttgat 320  
 tttgtcagtg ttttctacca tccctgagca cacaaaattg 360  
 gcctcaagtt gcctcttgat cctggagtgc gtgatgattg 400  
 tcgtcttttg tttggagtgc atcattcgaa tctggtctgc 440  
 gggttgctgt tgtcgatata gaggatggca aggaagactg 480  
 aggtttgctc gaaagccctt ctgtgttata gataccattg 520  
 ttcttatcgc ttcaatagca gttgtttctg caaaaactca 560  
 gggtaatatt tttgccacgt ctgcactcag aagtctccgt 600  
 ttcttacaga tcctccgcat ggtgcgcag gaccgaaggg 640  
 gaggcacttg gaaattactg gggtcagtgg tttatgctca 680  
 cagcaaggaa ttaatcacag cttggtacat aggatttttg 720  
 gttcttattt tttcgtcttt ccttgtctat ctggtggaaa 760

FIG. 1B

aggatgccaa	taaagagttt	tctacatatg	cagatgctct	800
ctggaggggc	acaattacat	tgacaactat	tggctatgga	840
gacaaaactc	ccctaacttg	gctgggaaga	ttgctttctg	880
caggctttgc	actccttggc	atttctttct	ttgcacttcc	920
tgccggcatt	cttggctcag	gttttgctcatt	aaaagtacaa	960
gaacaacacc	gccagaaaaca	ctttgagaaa	agaaggaacc	1000
cagctgccaa	cctcattcag	tgtgtttggc	gtagttacgc	1040
agctgatgag	aaatctgttt	ccattgcaac	ctggaagcca	1080
cacttgaagg	ccttgccacac	ctgcagccct	accaagaaaag	1120
aacaagggga	agcatcaagc	agtcagaagc	taagttttaa	1160
ggagcgagtg	cgcattggcta	gccccagggg	ccagagtatt	1200
aagagccgac	aagcctcagt	aggtgacagg	aggtccccaa	1240
gcaccgacat	cacagccgag	ggcagtccca	ccaaagtgca	1280
gaagagctgg	agcttcaacg	accgaaccgg	cttcggggcc	1320
tcgctgcgcc	tcaaaagttc	tcagccaaaa	ccagtgatag	1360
atgctgacac	agcccttggc	actgatgatg	tatatgatga	1400
aaaaggatgc	cagtgtgatg	tatcagtgga	agacctcacc	1440
ccaccactta	aaactgtcat	tcgagctatc	agaattatga	1480
aatttcattgt	tgcaaaacgg	aagtttaagg	aaacgttacg	1520
tccatatgat	gtaaaagatg	tcattgaaca	atattctgct	1560

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FIG. 1C

ggtcacatctgg	acatgtttgtg	tagaattaaa	agccttcaaa	1600
cacgtgttga	tcaaattctt	ggaaaagggc	aaatcacatc	1640
agataagaag	agccgagaga	aaataacagc	agaacatgag	1680
accacagacg	atctcagtat	gctcggtcgg	gtgggtcaagg	1720
ttgaaaaaca	ggtagagtcc	atagagtcca	agctgggactg	1760
cctactagac	atctatcaac	aggtccttcg	gaaaggctct	1800
gcctcagccc	tcgctttggc	ttcattccag	atcccacctt	1840
ttgaatgtga	acagacatct	gactatcaaa	gccctgtgga	1880
tagcaaagat	ctttcgggtt	ccgcacaaaa	cagtggctgc	1920
ttatccagat	caactagtgc	caacatctcg	agaggcctgc	1960
agttcattct	gacgccaaat	gagttcagtg	cccagacttt	2000
ctacgcgctt	agccctacta	tgcacagtca	agcaacacag	2040
gtgccaatga	gtcaaagcga	tggctcagca	gtggcagcca	2080
ccaacacccat	tgcaaaccac	ataaatacgg	cacccaagcc	2120
agcagcccca	acaactttac	agatcccacc	tcctctccca	2160
gccatcaagc	atctgcccag	gccagaaact	ctgcacccta	2200
accctgcagg	cttacaggaa	agcattttctg	acgtcaccac	2240
ctgccttggt	gcctccaagg	aaaatgttca	ggttgcacag	2280
tcaaatctca	ccaaggaccg	ttctatgagg	aaaagctttg	2320
acatggggagg	agaaactctg	ttgtctgtct	gtcccatggt	2360
gccgaaggac	ttggggcaaat	ctttgtctgt	gcaaaacctg	2400
atcaggtcga	ccgaggaact	gaatatacaa	ctttcaggga	2440

FIG. 1D

gtgagtcaag tggctccaga ggcagccaag atttttaccc 2480  
caaatggagg gaatccaaat tgtttataac tgatgaagag 2520  
gtgggtcccg aagagacaga gacagacact tttgatgccg 2560  
caccgcagcc tgccagggaa gctgcctttg catcagactc 2600  
tctaaggact ggaagggtcac gatcatctca ggcatttgt 2640  
aaggcaggag aaagtacaga tgccctcagc ttgcctcatg 2680  
tcaaactgaa ataa 2694

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FIG. 2A

KCNQ5 Protein Sequence

MKDVESGRGR	VLLNSAAARG	DGLLLLGTTRA	ATLGGGGGGL	40
RESRRGKQGA	RMSLLGKPLS	YTSSQSCRNR	VKYRRVQNYL	80
YNVLERPRGW	AFIYHAFVFL	LVFGCLILSV	FSTIPEHTKL	120
ASSCLLILEF	VMIVVFGLEF	IIRIWSAGCC	CRYRGWQGR	160
RFARKPFCVI	DTIVLIASIA	VVSAKTQGNI	FATSALRSLR	200
FLQILRMVRM	DRRGGTWKLL	GSVVYAHKE	LITAWYIGFL	240
VLIFSSFLVY	LVEKDANKEF	STYADALWWG	TITLTTIGYG	280
DKTPLTWLGR	LLSAGFALLG	ISFFALPAGI	LGSGFALKVQ	320
EQHRQKHFEK	RRNPAANLIQ	CVWRSYAADE	KSVSIATWKP	360
HLKALHTCSP	TKKEQGEASS	SQKLSFKERV	RMASPRGQSI	400
KSRQASVGDR	RSPSTDITAE	GSPTKVQKSW	SFNDRTFRFP	440
SLRLKSSQPK	PVIDADTALG	TDDVYDEKGC	QCDVSVEDLT	480
PPLKTVIRAI	RIMKFHVAKR	KFKETLRPYD	VKDVIQYSA	520
GHLDMLCRIK	SLQTRVDQIL	GKGQITSDDK	SREKITAHEE	560
TTDDLMLGR	VVKVEKQVQS	IESKLDCLLD	IYQQVLRKGS	600
ASALALASFQ	IPPFCEQTS	DYQSPVDSKD	LSGSAQNSGC	640
LSRSTSANIS	RGLQFILTPN	EFSAQTFYAL	SPTMHSQATQ	680
VPISQSDGSA	VAATNTIANQ	INTAPKPAAP	TTLQIPPLP	720
AIKHLPRPET	LHPNPAGLQE	SISDVTTCLV	ASKENVQVAQ	760
SNLTKDRSMR	KSFDMGGETL	LSVCPMVPKD	LKSLSVQNL	800
IRSTEELNIQ	LSGSESSGSR	GSQDFYPKWR	ESKLFITDEE	840

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FIG. 2B

VGPEETETDT FDAAPQPARE AAFASDSLRT GRSRSSQSIC 880

KAGESTDALS LPHVKLK 897

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104350-0209980

FIG. 3

**Alternative Splice Exon 1**

TGG	GGA	CAG	TGG	ACA	TTG	CGT
Trp	Gly	Gln	Trp	Thr	Leu	Arg

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# FIG. 4A

## 3' UTR

gttcttcatt	ttctttccag	gcatagcagt	tcttttagcca	40
tacatatcat	tgcatgaact	atttcgaaag	cccttctaaa	80
aagttgaaat	tgcaagaatc	gggaagaaca	tgaaaggcag	120
tttataagcc	cgttaccttt	taattgcatg	aaaatgcatg	160
tttagggatg	gctaaaatc	caagggtgat	cgacattaac	200
ccactcattt	agtaatgtac	cttgagttaa	aaagcctgag	240
aaaccaaaca	cagctaatac	tatgggggtg	atgaatatgt	280
caagtttagg	tcatttagaa	gatttgacac	tgtattttga	320
aattatgagt	aaacaccttc	aaatttcagg	cattttctgt	360
ttgtgactaa	atacaaaact	cattttcaag	attaggccat	400
aatgtatat	taaacacaat	ggctatcaac	agctgctaata	440
aagggtatcaa	ctaaagcaga	attgggggaat	aatagaaatg	480
gctgcttatt	tcaagatata	tttgccaacc	cattcctatt	520
cagtcatttt	attattaatg	taatttgaat	gtcaatttgt	560
gtgcttttgg	tgatttagcg	ctgtggcaag	caattttgca	600
catcattttc	atgttggtct	ttatgacaag	aatgtttctc	640
aattagaaaa	tgtgcaaata	atgaaattca	gggccagtga	680
ggcaaataga	ctatctgaca	tatttgactt	tatgaaaaca	720
tattgcctga	tggcagaatc	aactttataa	gtggccaact	760
tctacacaag	cgtatgaaat	actggtcagt	agaacagcca	800



FIG. 4B

ttgtgattgg actggtttct ctgcaatggc gccaaaccca	840
ggcttgccaa tactgcctat gtaaaggga agtgtgagaa	880
gctatttctca ttctgctgac atacaggtag gactatgggg	920
gatggggacat ttgagtggga ctgagatagg aaaggcttga	960
aaagaaccca gaaacaccac caggaagttg gcaaagtaaa	1000
agaaaatgac ttccccctca aagggaatg agagggagag	1040
aaacaaacca aaatagaaga actagacttt ttagaaaatg	1080
agtattgcta	1090

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FIG. 5A

Multiple Sequence Alignment of the KCNQ Channel Family Members

hskcnq4 ~~~~~MABAPPRRLGLGPPPGDAPRAE..LVALTAVQSEQCEEG.  
hskcnq5 ~~~~~MKDVESGR..GRVLLNSAAARGDGLLLGLTRAAATLGGGG.  
hskcnq2 ~~~~~MVQKSRNG..GVYPG..PSEGEKLLKVGFGVGLDGP..AP.  
hskcnq3 MGLKARRAAGAAGGGGDDGGGGGGAANPAGGDAAGDEERKVGGLAPGDVEQVTLALGA.  
hskcnq1 ~~~~~MAAASSPPRAERKRKWGRLPGARRGSAGLAKKCPFSLELAEQGP

hskcnq4 GGGSPRRLGLLGLSPLP..PGAPLPGP GSGSACGGRSSAA..HKR...Y..RR.....  
hskcnq5 GGLRESRRGKQCARMSLLGKPL...SYT..SSQSC..RR...NVK...Y..RR.....  
hskcnq2 DSTRDGALLIACSEAPKRGSI LSKPRAG..GAGA...CKPPKR..NAF...Y..RK.....  
hskcnq3 GADKDGTLLEGGGRDE..GQRRTPGGIGLLAKTFLSPVKRRNAK...Y..RR.....  
hskcnq1 AGGALYAPIAPCAPGPAPPASPAPAAPPVASDLGPFPVSLDPRVSTSTRRPVLARTH

hskcnq4 LONWYVNVLERPRGW..AFYVYHVFVFLVVFSCVLSLSLTIQEHQELANECILIEFVMIV  
hskcnq5 VONYLYNVLERPRGW..AFYVYHAFVFLVVFGLLSVFSFIPEHTKLASSCLLIEFVMIV  
hskcnq2 LCNFLYVNVLERPRGW..AFYVYHAFVFLVVFSCVLSVFSFIKEYEKSSEGAIVILBIVTIV  
hskcnq3 TCTLTYDALERPRGW..ALFYHALVFVFLVGLCLLAWLTIKFKEYTVSGDWMLDLBTATF  
hskcnq1 MGRMYNFLERPTFWKCFVYHFAFVFLVFLVLSLTIQEQYALATGTFTFWMEIVLVV

hskcnq4 VFGLEYIVRWWSGCCCRVRCWGRFARFKPFVVDITIVFVASVAVLAAGTQCNIFATS  
hskcnq5 VFGLEYIVRWWSGCCCRVRCWGRFARFKPFVVDITIVFVASVAVSAKTQCNIFATS  
hskcnq2 VFGVEYIVRWWSGCCCRVRCWGRFARFKPFVVDITIVFVASVAVLAAGSQCNIFATS  
hskcnq3 IFGADFAIRWWSGCCCRVRCWGRFARFKPLCMIDITIVFVASVAVPVAAGNQCNIATLS  
hskcnq1 FFGLEYIVRWWSGCCCRVRCWGRFARFKPLISITIDITIVFVASVAVVGCVSKQCNIFATS

hskcnq4 ALRSRLFLQILRMVRDRRGGTWKL LGSVVYAHSKELTAWYIGFLVLIFASRLVYLAER  
hskcnq5 ALRSRLFLQILRMVRDRRGGTWKL LGSVVYAHSKELTAWYIGFLVLIFASRLVYLVVEK  
hskcnq2 ALRSRLFLQILRMVRDRRGGTWKL LGSVVYAHSKELTAWYIGFLVLIFASRLVYLAER  
hskcnq3 ALRSRLFLQILRMVRDRRGGTWKL LGSVAFAHSKELTAWYIGFLVLIFASRLVYLVVEK  
hskcnq1 ALRSRLFLQILRMVRDRRGGTWKL LGSVVYAHSKELTAWYIGFLVLIFASRLVYLAER

hskcnq4 DA.....NSDFSSYADSLWNGTITTTTIGYGDKPHITWLGRVLAAGFAELGHSFF  
hskcnq5 DA.....NKEFSTYADALWNGTITTTTIGYGDKPHITWLGRVLAAGFAELGHSFF  
hskcnq2 GE.....NDHEDTYADALWNGTITTTTIGYGDKPHITWLGRVLAAGFAELGHSFF  
hskcnq3 DVPEVDAQGEEMKEEFYADALWNGTITTTTIGYGDKPHITWLGRVLAAGFAELGHSFF  
hskcnq1 DAVNESGRV.....EFGSYADALWNGTITTTTIGYGDKPHITWLGRVLAAGFAELGHSFF

hskcnq4 ALPAGILGSGFLKVOEQHROKHFEKRRMPAANLIQAARLSTDMRSRAYLTATWYYYS  
hskcnq5 ALPAGILGSGFLKVOEQHROKHFEKRRMPAANLIQAARLSTDMRSRAYLTATWYYYS  
hskcnq2 ALPAGILGSGFLKVOEQHROKHFEKRRMPAANLIQAARLSTDMRSRAYLTATWYYYS  
hskcnq3 ALPAGILGSGFLKVOEQHROKHFEKRRMPAANLIQAARLSTDMRSRAYLTATWYYYS  
hskcnq1 ALPAGILGSGFLKVOEQHROKHFEKRRMPAANLIQAARLSTDMRSRAYLTATWYYYS

hskcnq4 ..ILPSFRELALLFEHVQARNGGLRPLEVRAPVPDGPASRYPPVATCHRPGSTSPCPG  
hskcnq5 ...KPHLKA.....HT.....CSPTK.....KEQGEAS....  
hskcnq2 TVTVPMYRLIPLPNQLELLRNLSKSGLAFRKDPPEPSP.....  
hskcnq3 VVSFPFPR.....KEQLEAA....  
hskcnq1 .....IRKAP..RSHTLLS.....PSPKPKSVVVKKKFKLKDNGVT

FIG. 5B

hskcnq4 ESSRMGIKDRIRMSSQRRRTGPS.KQQLAPPTMPTSPSSSEQVGEATSETKVQKSWSFNDR  
 hskcnq5 SSQKHSFKERYRMASPRGQSIKS.RQ..ASVGDRRSESTDITAEG.SP TKVQKSWSFNDR  
 hskcnq2 .SQKLSLKDRY.FSSPRGVAAG.KGSPQAQTVRRSEADQSLE.DSPSKVPKSWSPGDR  
 hskcnq3 .SSQKLGLLDRYFSSNPRGSNTKG.KLF.....DPLNVDAIE.ESSPKPKPVGLNNK  
 hskcnq1 PGEKMLTVPHHTCDPPERRLDHFSVDGYDSVRKSEPT...LLEWSMP.....

hskcnq4 TRFRASLRL....KPRTSADA.PSEEVAEKSYQCELTVDIMPAVITVTRSTRIEKE  
 hskcnq5 TRFRPSLRRLKSSQPKFVIDASTALGTTDDVYDEKGCQCDVSVEDLTPPLKTVTRATRIKMF  
 hskcnq2 SRARQAFRIKGAASR.QNSEEASLPGEDIVDDKSCPCFVTEDETPGLKVSIRAVCMVRP  
 hskcnq3 ERFRTAFRMKAYAFWQ..SSEDAGTGDPMADRGYGNDFIEDMIPTLKAAHRAVRIHQF  
 hskcnq1 .HF...MTNSFAEDLDLEGETLLT..PETH.....ISQREHHRATKVRRMQY

hskcnq4 LYAKKKFKETLRLPYDVMDVIEQYSAGHLDMESRIKSLQTRVQIVVGRG...PGDR.KARE  
 hskcnq5 HVAKKFKETLRLPYDVMDVIEQYSAGHLDMESRIKSLQTRVQIVVGRG...PGDR.KARE  
 hskcnq2 LUSKKRKFESLRLPYDVMDVIEQYSAGHLDMESRIKSLQTRVQIVVGRGPA.ITD..KDR.  
 hskcnq3 RLYKKFKETLRLPYDVMDVIEQYSAGHLDMESRIKSLQTRVQIVVGRGPA.ITD..KDR.  
 hskcnq1 FVAKKKFQARKPYDVMDVIEQYSAGHLDMESRIKSLQTRVQIVVGRGPA.ITD..KDR.

hskcnq4 KGDKG.....PSDAEVVDEISMGRRVVKVE..KQVQSIEHKLCLLLGFY  
 hskcnq5 ...KI.....TAHEHTTDDLSMLGRVVKVE..KQVQSIEHKLCLLLDIY  
 hskcnq2 ..TKG.....PAAELPEDPSMGLRGVKE..KQVLSMEKKDLFLVNIY  
 hskcnq3 KGSFTFPSSQSPRNEPYVARPSTSEI.EDQSMGKFVKVE..RQVQDMGKKDLFLVDMH  
 hskcnq1 RGSNTIGARLNVRVEDKVTQLDQRLALITD...MEHQLLSEHGGSTPGSGGPPREGGAHIT

hskcnq4 SRCLRSRG...SA.SLGAVQVPLFPDITSDYHSPVTH..EDISVSAQTLIS.ISRSVSTNM  
 hskcnq5 QQVLRKGSA.SALALASFQIPPFCEQTSYQSPVBS..KDLSGSAQNSGCLSRSTSANI  
 hskcnq2 MQ..RMGIP.PTETEYFGAK..PEPAPPYHSPEDS..RE...HVDRHGCIKIVRSSS  
 hskcnq3 MQHMER.....LQVQVTEYPTKGTSSPAAEKKEDNRYSLDKTIICNYSETGP  
 hskcnq1 QPCSGSGSVDPFLFLPSNTLPTYE.QLTVPRRGEDRGS~~~~~

hskcnq4 D~~~~~  
 hskcnq5 SRGLQFI..LTPNEFSAQTFYALSPTMHSAQATQVPIQSQSDGSAVAATNTIANQINTAPKP  
 hskcnq2 STGQKNF..SAPPAAPP...VQCPPTSWQPSHPRQGHGTSVGDHGLSVIRIPPPAH  
 hskcnq3 PEPYSPHQVTIDKVSFYGFFAHDVFNLRGSGPSS.GKVQATPPSSATTYVERPTVLPIL  
 hskcnq1 ~~~~~

hskcnq4 ~~~~~  
 hskcnq5 AAPTTLQIPPLPAIKHLPRPETLHPNPAQLQESISDVTTCLVASKENVQVAQSNLTKDR  
 hskcnq2 ERLSAYGGGNRASMEFLRQEDTPGCRPPEGTLRDSITSISIPSVDHHEELERSPSGFSIS  
 hskcnq3 TLLDSRVSCSHQADLQG.PYSDRISFRQRRSITRDSDTPLSLMSVNHHEELERSPSGFSIS  
 hskcnq1 ~~~~~

hskcnq4 ~~~~~  
 hskcnq5 SMRKSFDMGGETLLSVCPMPVK...DLGKSLSVQNLIRSTEELNIQSGSESSESGRSGQ  
 hskcnq2 QSKENLDALNCSYAAVAPCAKVRPYIAEGESDTSDDLCTPCGPPPRSATGEGPFGDVWGA  
 hskcnq3 QDRDDYVFGPN...GGSSWMREKRYLAEGETDTPDFTPPSGSMPLSSTGDGIDSVDVWTP  
 hskcnq1 ~~~~~

FIG. 5C

```

hskcnq4 ~~~~~
hskcnq5 DFYPKWRESKLFITDEEVGPEETETDTFDAAPQPAREAAFASDSLRTGRSRSSQSICKAG
hskcnq2 GPRK~
hskcnq3 SNKPI~
hskcnq1 ~~~~~

hskcnq4 ~~~~~
hskcnq5 ESTDALSLPHVKLK
hskcnq2 ~~~~~
hskcnq3 ~~~~~
hskcnq1 ~~~~~

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FIG. 6A

## Human RNA Master Blot

TABLE 1

	1	2	3	4	5	6	7	8
A	whole brain	amygdala	Caudate nucleus	cerebellum	cerebral cortex	frontal lobe	hippocampus	medulla oblongata
B	occipital pole	putamen	substantia nigra	temporal lobe	thalamus	Subthalamic nucleus	spinal cord	
C	heart	aorta	Skeletal muscle	colon	bladder	uterus	prostate	stomach
D	testis	ovary	pancreas	pituitary gland	adrenal gland	thyroid gland	salivary gland	mammary gland
E	kidney	liver	small intestine	spleen	thymus	peripheral leukocyte	lymph node	bone marrow
F	Appendix	lung	trachea	placenta				
G	fetal brain	fetal heart	fetal kidney	fetal liver	fetal spleen	fetal thymus	fetal lung	
H	yeast total RNA	yeast tRNA	<i>E. coli</i> rRNA	<i>E. coli</i> DNA	Poly r(A)	human C <sub>0</sub> t DNA	human DNA	human DNA

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FIG. 6B

	1	2	3	4	5	6	7	8
A	•		•		*	*	•	
B	•	•		*				
C			*					
D								
E								
F								
G								
H				•				

FIG. 7

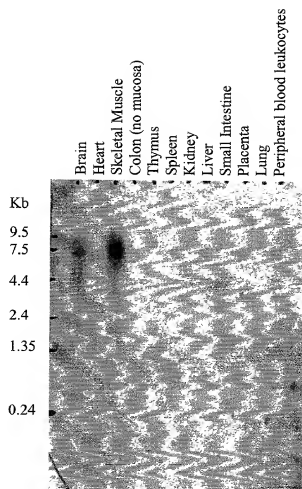
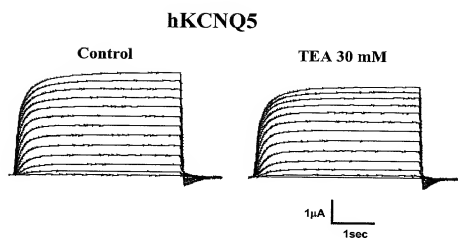


FIG. 8





**Antisense**



**FIG. 9A**



**FIG. 9B**



**FIG. 9C**

**Sense**



**FIG. 9D**



**FIG. 9E**



**FIG. 9F**

FIG. 10A

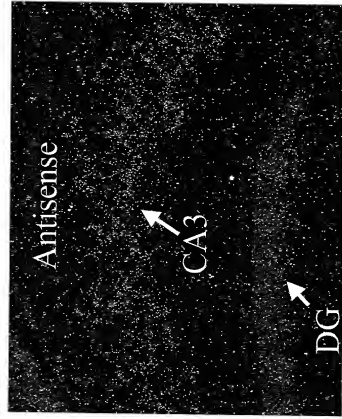


FIG. 10B

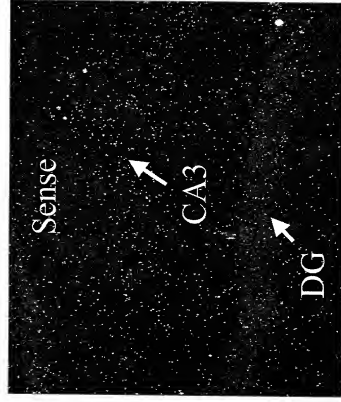


FIG. 11

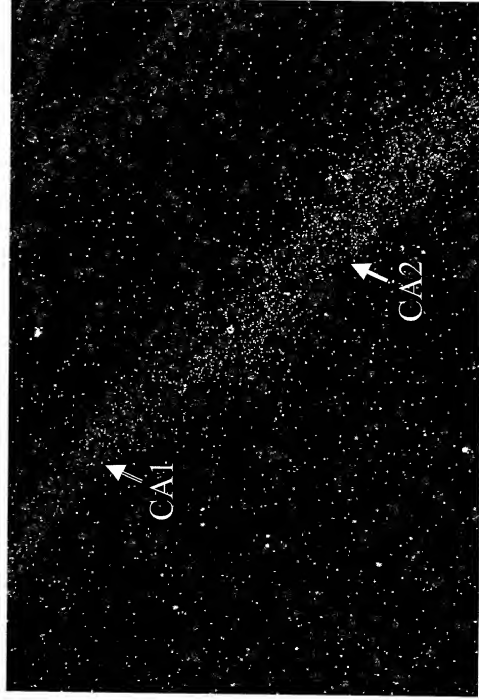


FIG. 12A

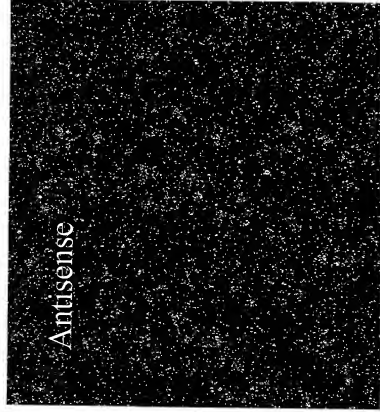


FIG. 12B

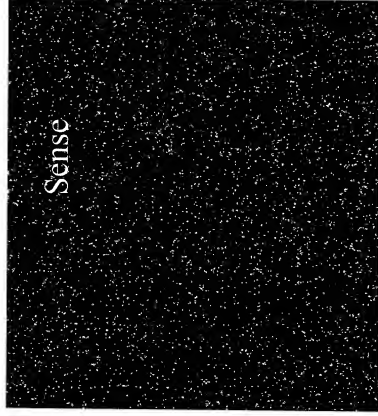


FIG. 13A

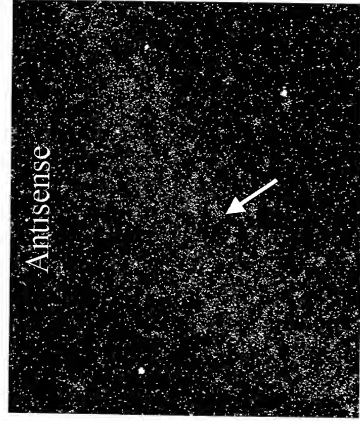


FIG. 13B

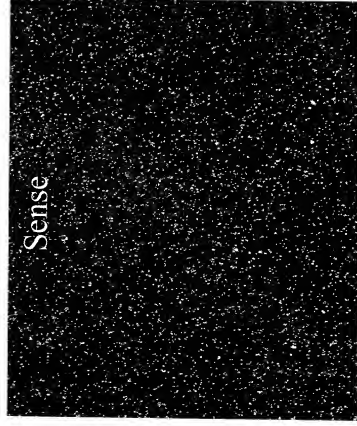


FIG. 14A

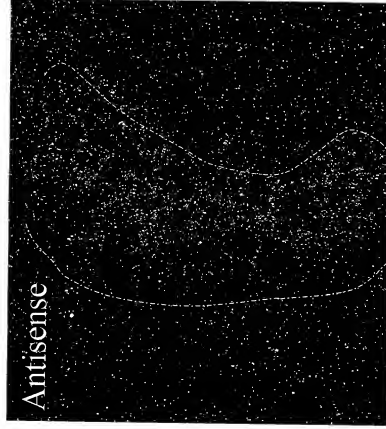


FIG. 14B

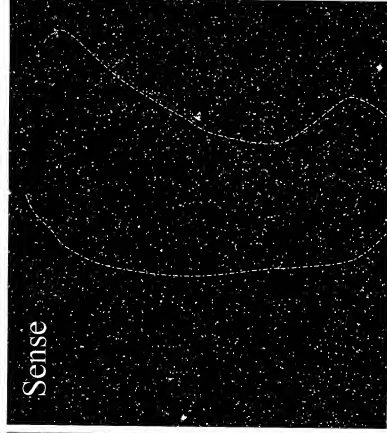


FIG. 15A

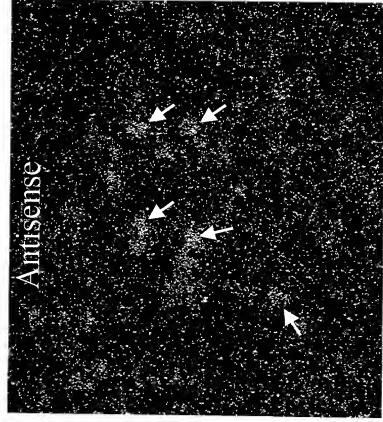


FIG. 15B

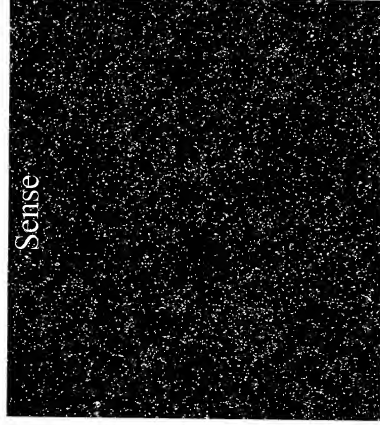


FIG. 16A

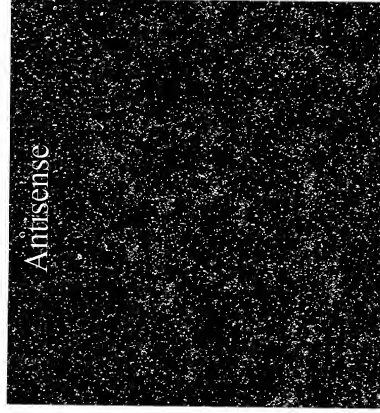


FIG. 16B

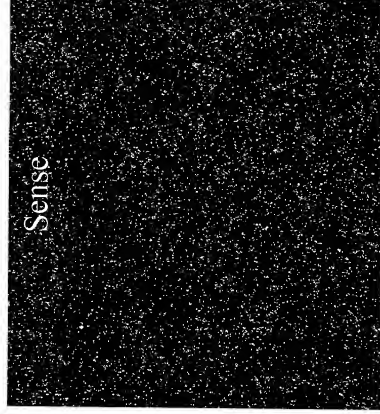




FIG. 17A

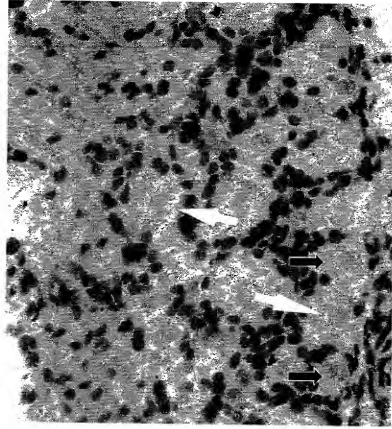


FIG. 17B

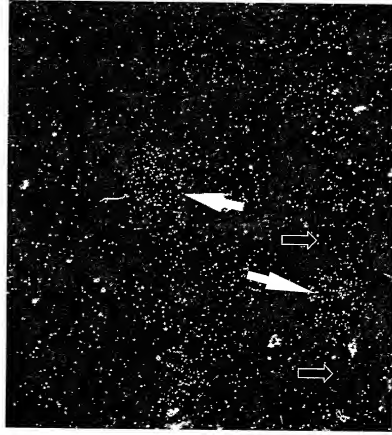


FIG. 18A

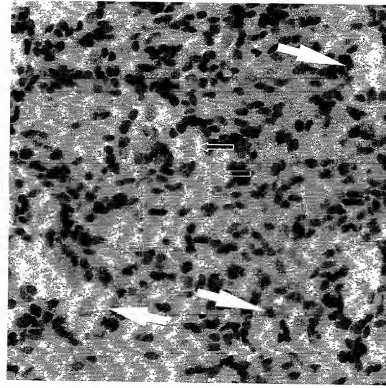


FIG. 18B

